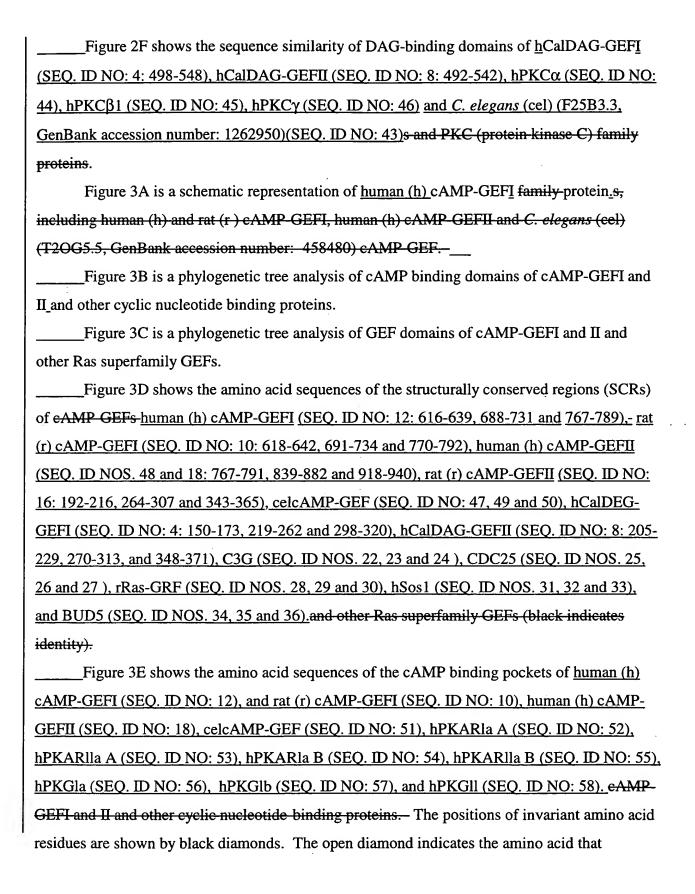
Amendments to the Specification

Please replace the Brief Description of the Drawings section, beginning on page 12 of the specification, with the following amended Brief Description of the Drawings section:

Brief Description of the Drawings

Figure 1 is a partial schematic diagram of a Ras pathway.

Figure 2A shows human (h) and mouse (m) CalDAG-GEFI, human (h) and rat (r) CalDAG-GEFII, and C. elegans (cel) (F25B3.3, GenBank accession number: 1262950) CalDAG-GEF. Figure 2B shows a computer-generated phylogenetic tree analysis of the GEF domains of hCalDAG-GEFI and hCalDAG-GEFII in relation to mCdc25, hSos1, C3G, rRas-GRF, C. elegans (cel) (F25B3.3, GenBank accession number: 1262950) and BUD5other-Ras superfamily GEFs. Figure 2C shows multiple alignment of GEF structurally conserved regions (SCRs) of F25B3.3 (SEQ. ID NOS. 19, 20 and 21), hCalDAG-GEFI (SEQ. ID NO: 4: 150-173, 219-262 and 298-320), hCalDAG-GEFII (SEQ. ID NO: 8: 205-229, 270-313, and 348-371), C3G (SEQ. ID NOS. 22, 23 and 24), mCdc25 (SEQ. ID NOS. 25, 26 and 27), rRas-GRF (SEQ. ID NOS. 28, 29 and 30), hSos1 (SEQ. ID NOS. 31, 32 and 33), BUD5 (SEQ. ID NOS. 34, 35 and 36) and C. elegans (cel) (F25B3.3, GenBank accession number: 1262950)(SEQ.ID NO: 37). s and several other GEFs of the Ras superfamily. Figure 2D shows the full-length amino acid sequences of human (h) (SEQ. ID NO: 4), and mouse (m) (SEQ. ID NO: 2) CalDAG-GEFI (box indicates amino acid differences). Figure 2E shows the sequence similarity (black indicates identity) of EF-hand domains in hCalDAG-GEFI (SEQ. ID NO: 4: 432-452), hCalDAG-GEFII (SEQ. ID NO: 8: 427-447), hCalmodulin (SEQ. ID NO: 38), hCalbindin D28K (SEQ. ID NO: 39), hCalcineurin B (SEQ. ID NO: 40), hParvalbumin α (SEQ. ID NO: 41), hTroponin C (SEQ. ID NO: 42), and C. elegans (cel)(F25B3.3, GenBank accession number: 1262950 (SEQ. ID NO: 37)s and other calcium binding proteins.



following amended row:

determines the binding specificity for cAMP or cGMP. The arrow indicates the position of amino acid substitutions specific to cAMP-GEFs.

______Figure 3F is the full-length amino acid sequences of human cAMP-GEFI (SEQ. ID NO: 12) and human cAMP-GEFII (SEQ. ID NO: 18)(boxes indicate amino acid identity).

On page 16 of the specification, line 1, please replace SEQ ID NO: 18 with SEQ ID NO: 17.

On page 16 of the specification, line 2, please replace SEQ ID NO: 19 with SEQ ID NO: 18.

On page 22, TABLE I, please replace the row beginning with hcAMP-GEFI with the